



New Knowledge – Fiber Diameter

- Fibers with a diameter of 0.4 µm are critical based on rat data
- Epidemiologic data suggest that fibers with diameter of 0.5 µm to 0.7 µm can reach the respiratory zone of the lungs
- There is some indication from epidemiologic data that fibers with a diameter as high as 1.5 µm can reach the respiratory zone of the lungs in mouth breathers
- Thus, fibers with a diameter of <0.5 μm to 1.5 μm are considered to be relevant as they can reach the respiratory zone of the lungs in humans









New Knowledge – Fiber Length

- Inhalation of fibers longer than 10 µm presents a considerably greater risk for lung cancer but the exact size cut-off for the length and magnitude of relative potency is uncertain
- There are two schools of thought about cancer toxicity of fibers <5 µm in length:
 - present a very low risk, possibly zero for cancer based on human data (?)
 - cause inflammation and may potentiate the pulmonary reaction to long fibers based on animal data and in vitro studies
- Fibers < 5 µm in length are causally associated with asbestosis and pleural plaques
- For mesothelioma, greater weight should be assigned to thinner fibers and fibers in the 5 µm to 10 µm in length range



New Knowledge – Fiber Type Lung Cancer

- There are different opinions about the relative potency of chrysotile vs amphiboles for lung cancer based on epidemiologic data
- Some assert that amphiboles are 5 times more toxic than chrysotile for lung cancer
- Others assert that no real difference is observed in statistical analysis of epidemiologic data
- The additional review of epidemiologic data to identify other factors such as industry in which exposure occurred might shed some light









New Knowledge – Fiber Type Mesothelioma

- It is becoming apparent that there are different relative carcinogenic potencies for different fiber types
- The available epidemiologic data provides compelling evidence that potency of amphiboles is at least two orders of magnitude greater than that of chrysotile (ATSDR)
- Time since first exposure is an important factor for occurrence of mesothelioma
- Duration and intensity of exposure is also found to be important in epidemiologic studies









New Knowledge -Other Amphibole Fibers

- Currently there are no data available either in humans or in animals about the toxicity of other amphiboles (e. g., winchite and richterite)
- Other amphibole fibers with similar durability and dimension would be expected to result in similar pathogenicity
- Thus, it may be prudent to consider potency of currently regulated and unregulated amphiboles to be similar









New Knowledge – Cleavage Fragments

- Data indicate that durability and dimension are critical to pulmonary pathogenesis
- There are little data directly addressing similarities and dissimilarities between the original fibers and cleavage fragments for pulmonary pathogenesis
- Evidence suggests that it is prudent at this time to assume equivalent potency for lung cancer in the absence of other information
- Similarly, evidence implies that for mesothelioma, thin diameter fibers and fibers >5 µm in length are found to be more important, thus, cleavage fragments that do not meet these criteria are not expected to contribute to the risk of mesothelioma





